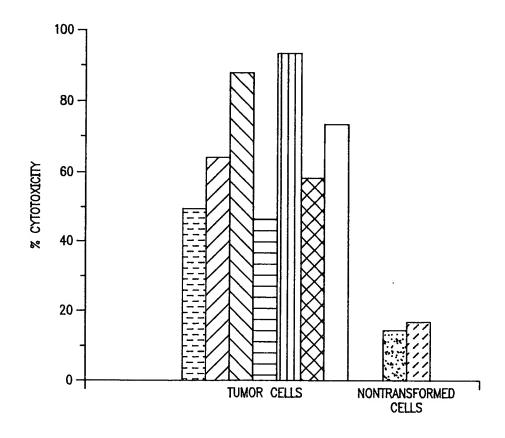


FIG. I





国 OCC1

A HEA

☐ OVCAR-3

JURKAT

LANCAP

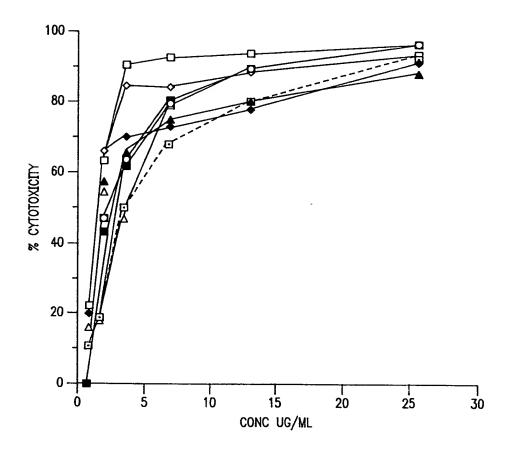
☐ FTC

☑ PS

HUVEC

FIG. 2





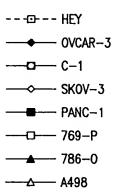
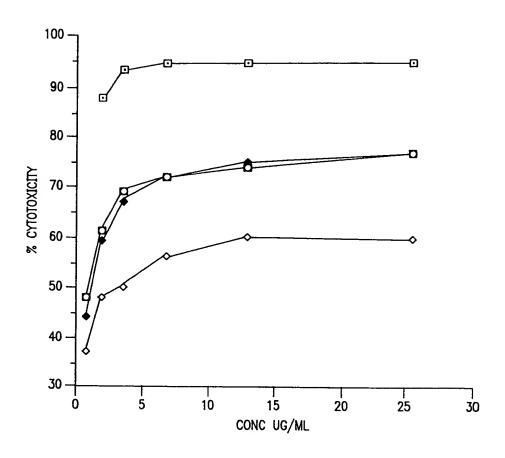


FIG. 3





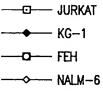
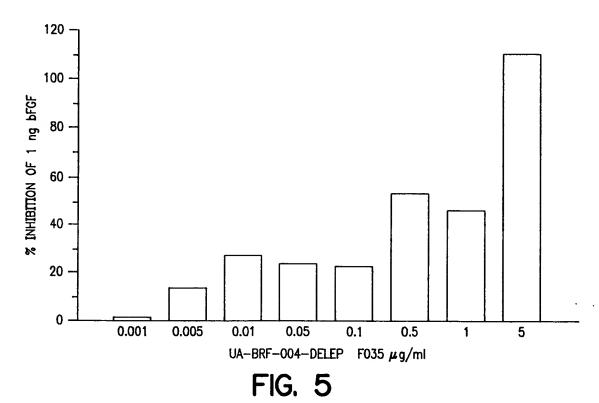


FIG. 4





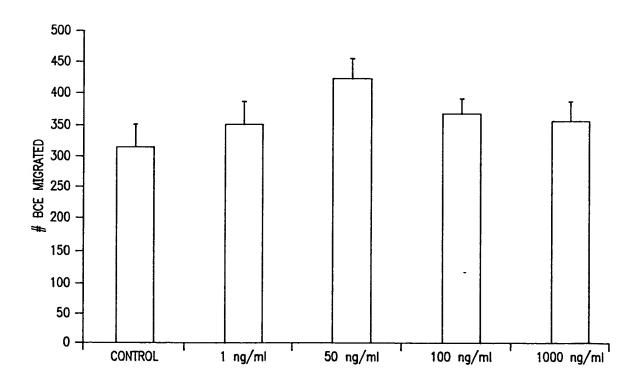


FIG. 6



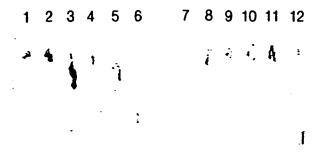


FIG. 7

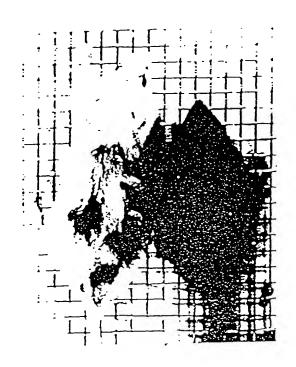


FIG. 8



Dose: 0.2 ml

ACETONE 4 Weeks

Mouse # 133



FIG. 9A

Dose: 0.2 ml

ACETONE 8 Weeks

Mouse # 150



FIG. 9B



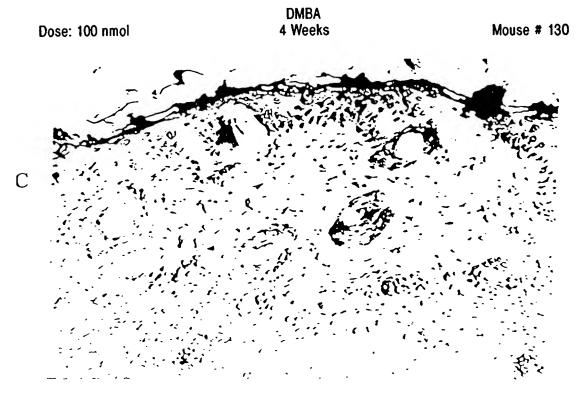


FIG. 9C

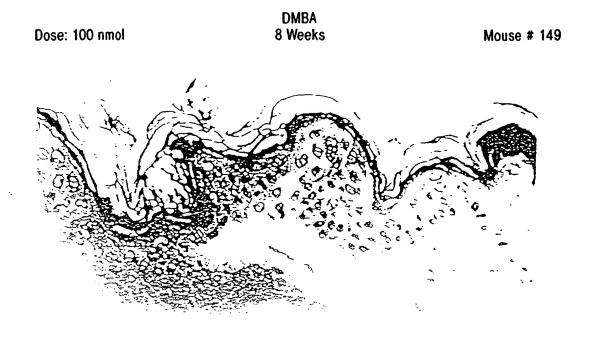


FIG. 9D



Dose: 0.3 mg

Compound # 35 4 Weeks

Mouse # 136



FIG. 9E

Dose: 0.3 mg

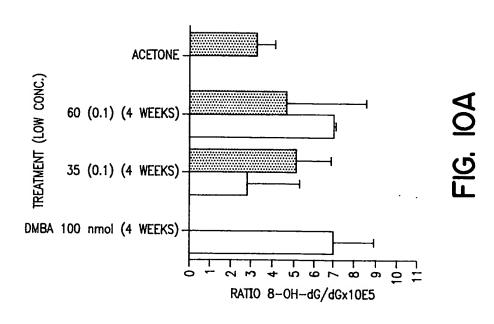
Compound # 35 + DMBA 8 Weeks

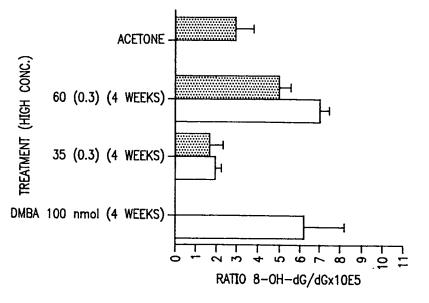
Mouse # 140



FIG. 9F



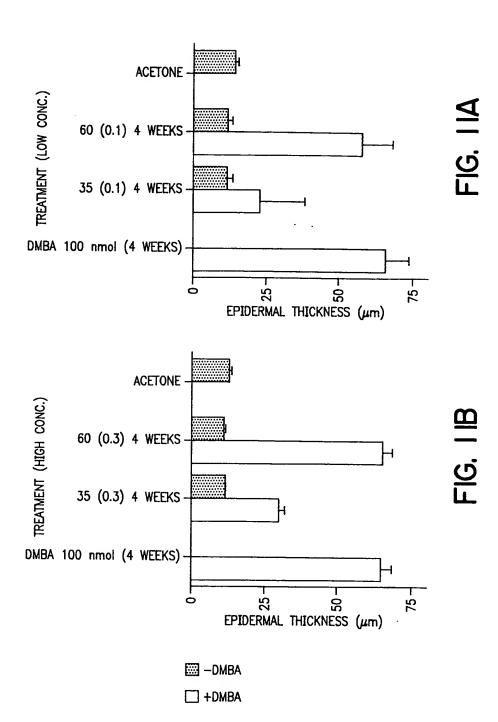




₩ -DMBA

-+DMBA







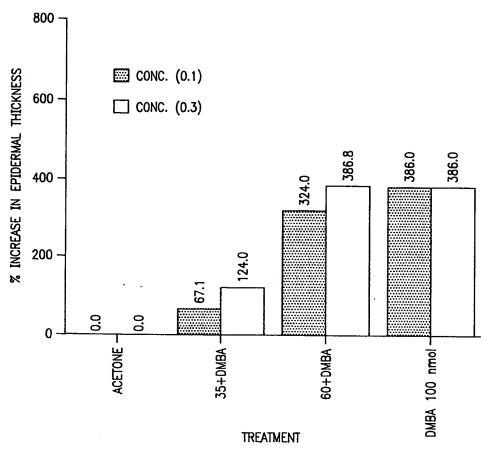


FIG. 12

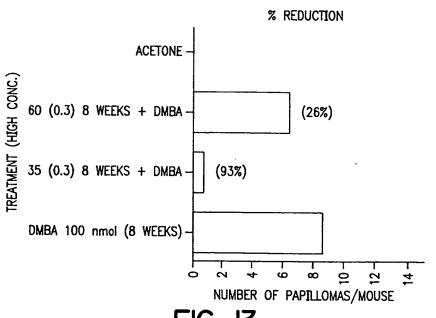


FIG. 13



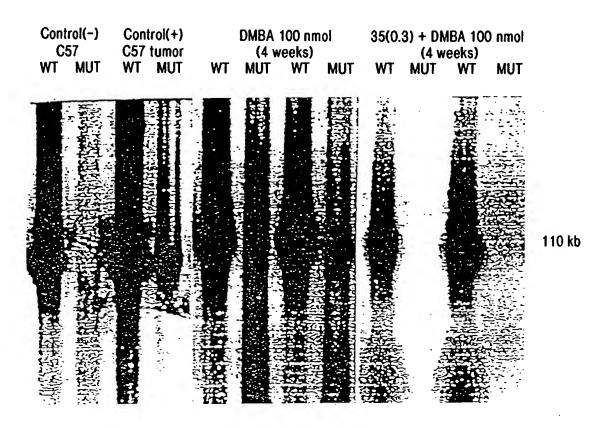


FIG. 14



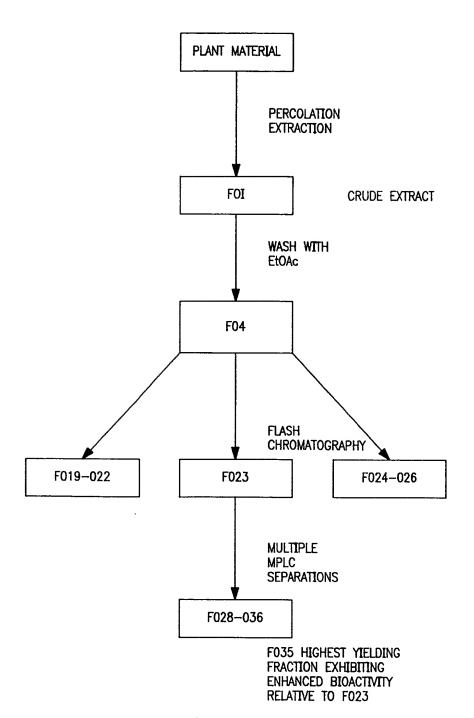


FIG. 15



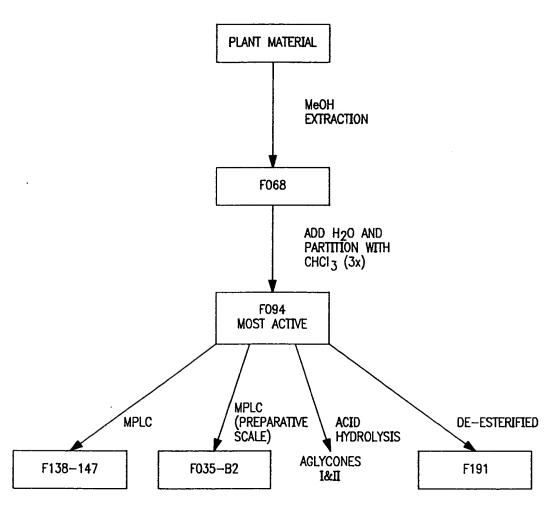
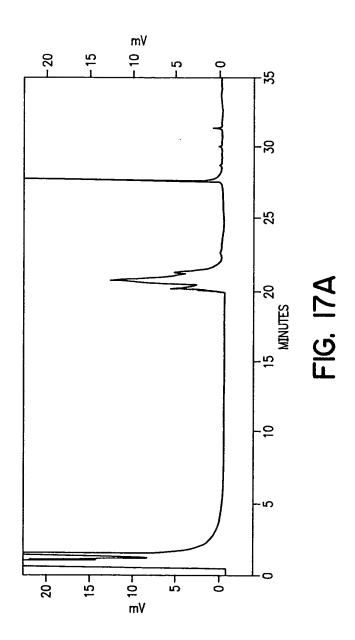
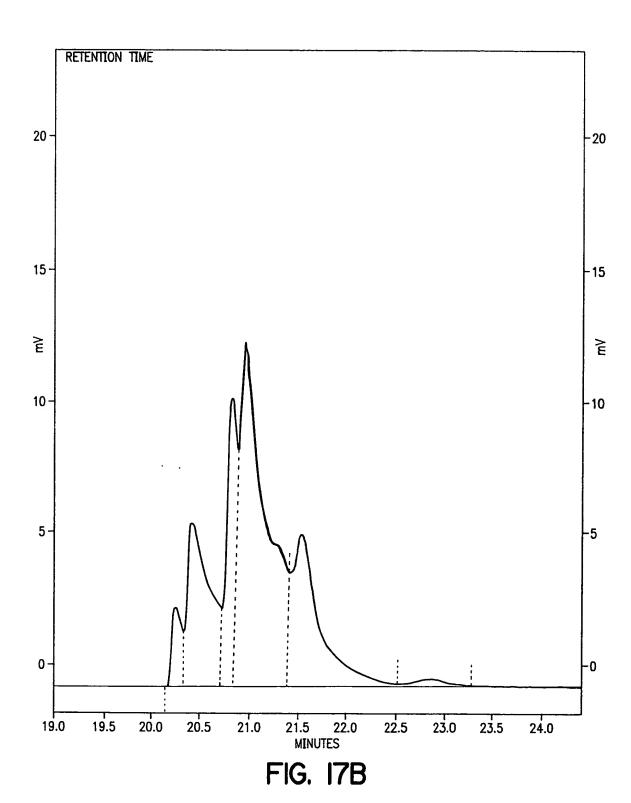


FIG. 16











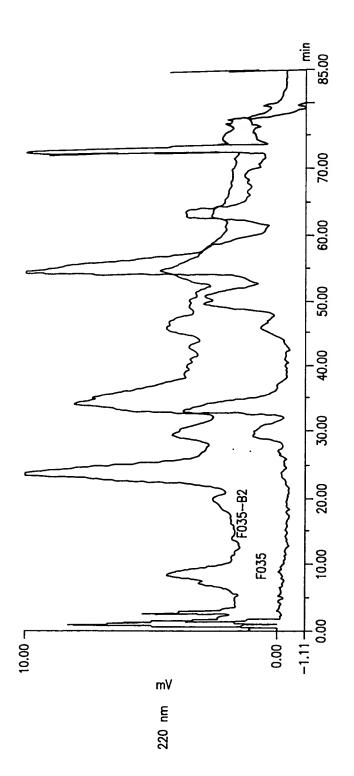
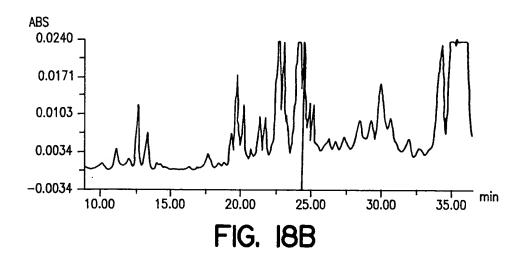
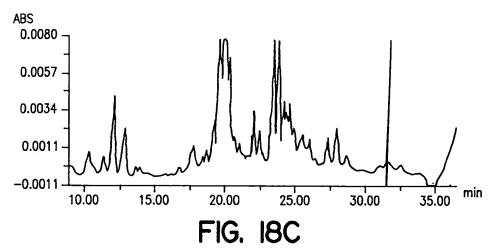
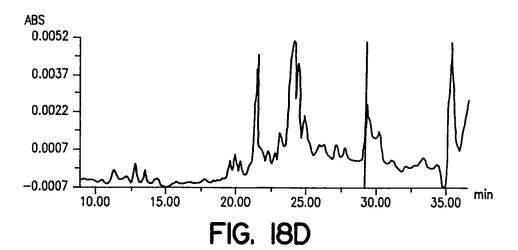


FIG. 18A

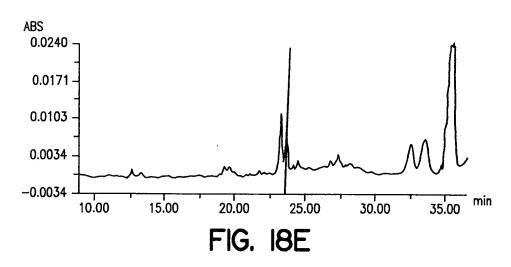


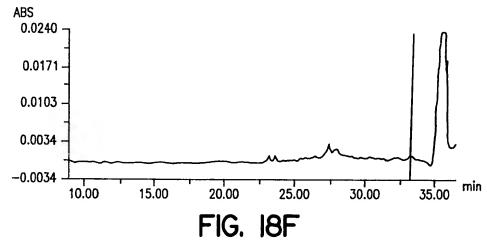




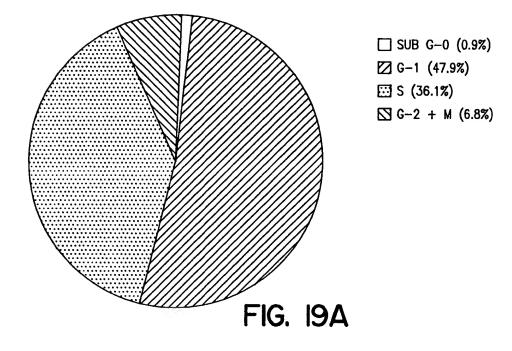


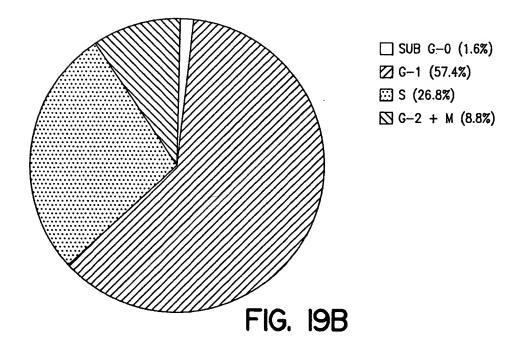




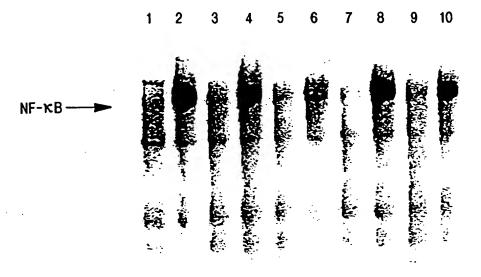












1:Untreated 2:TNF (100pM) 3:F035 (1µg/ml) 4:TNF + F035 (1µg/ml) 5:F035 (2µg/ml) 6:TNF + F035 (2µg/ml) 7:F094 (1µg/ml) 8:TNF + F094 (1µg/ml) 9:F094 (2µg/ml) 10:TNF + F094 (2µg/ml)

FIG. 20



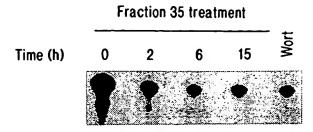


FIG. 21



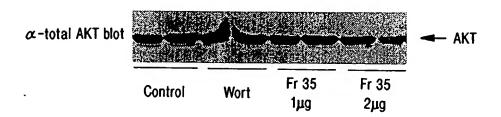


FIG. 22



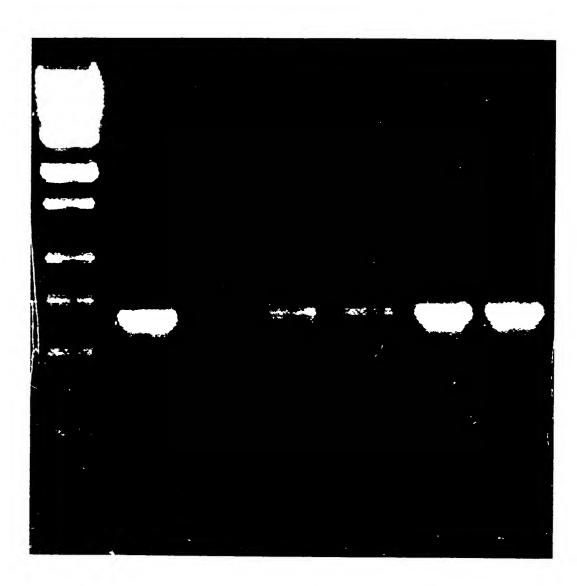


FIG. 23



STRUCTURE OF ELLIPTOSIDES:

ELLIPTOSIDE E IF R=OH

ELLIPTOSIDE A IF R=H



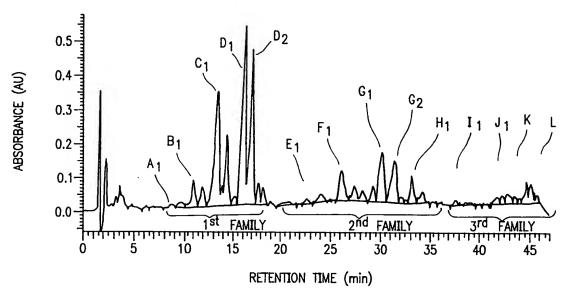


FIG. 25

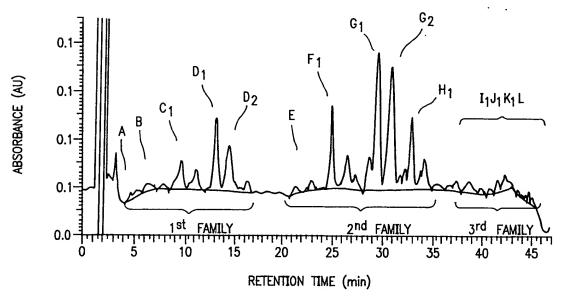
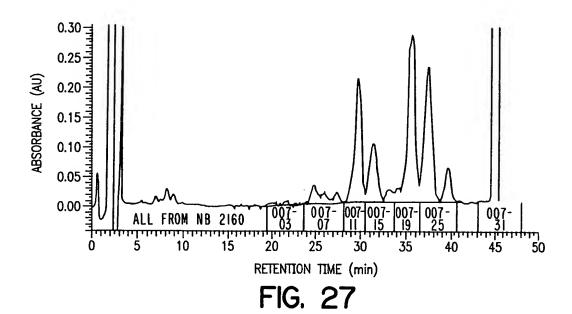
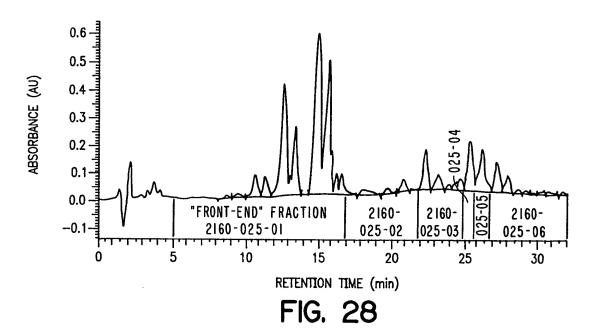


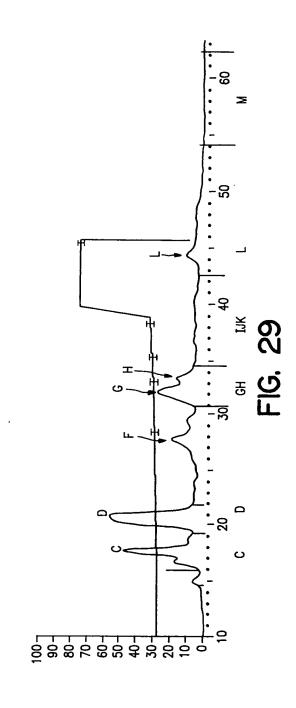
FIG. 26



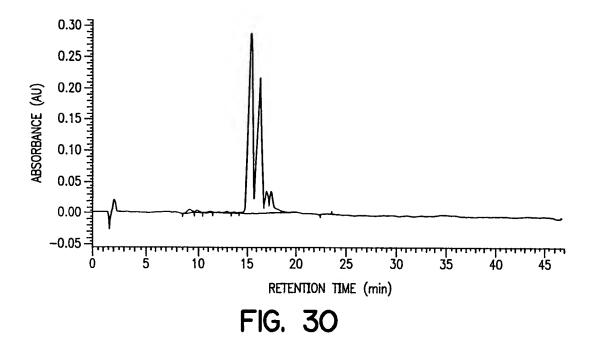












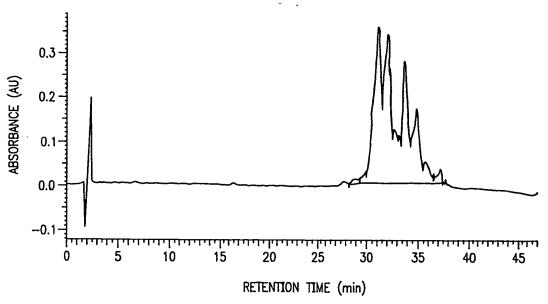
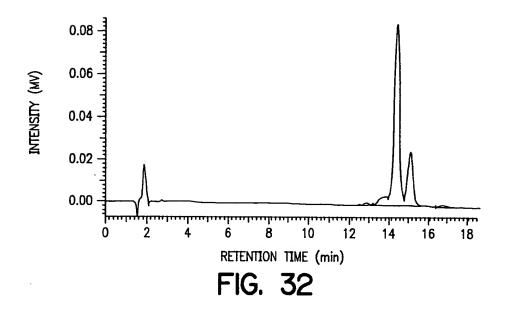
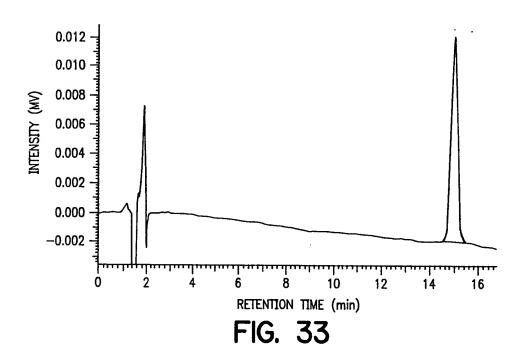


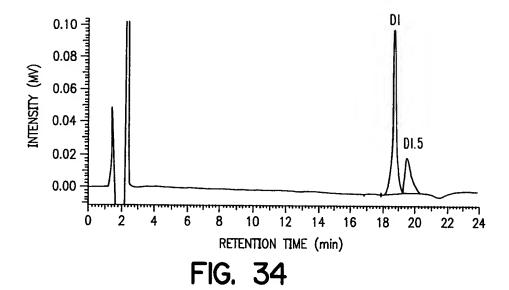
FIG. 31

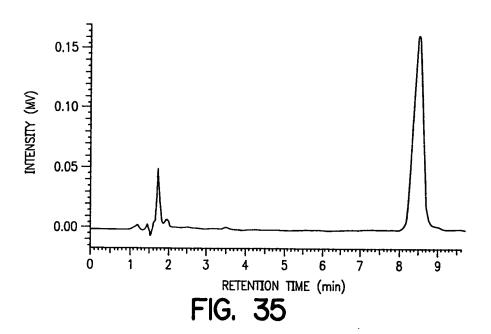




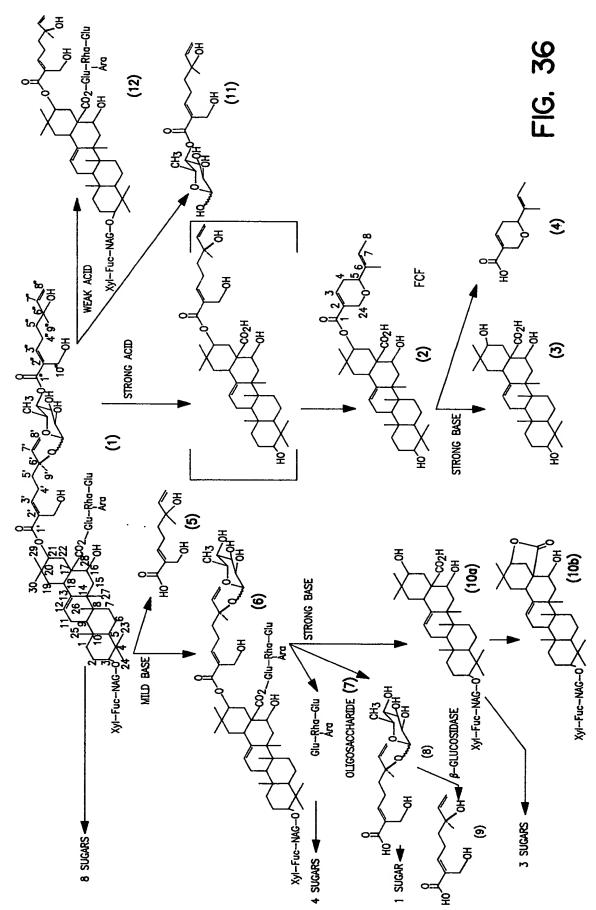


















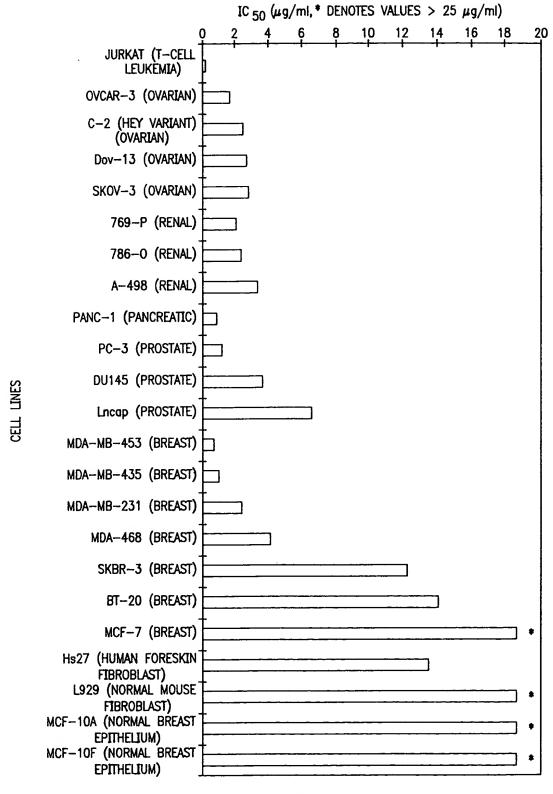
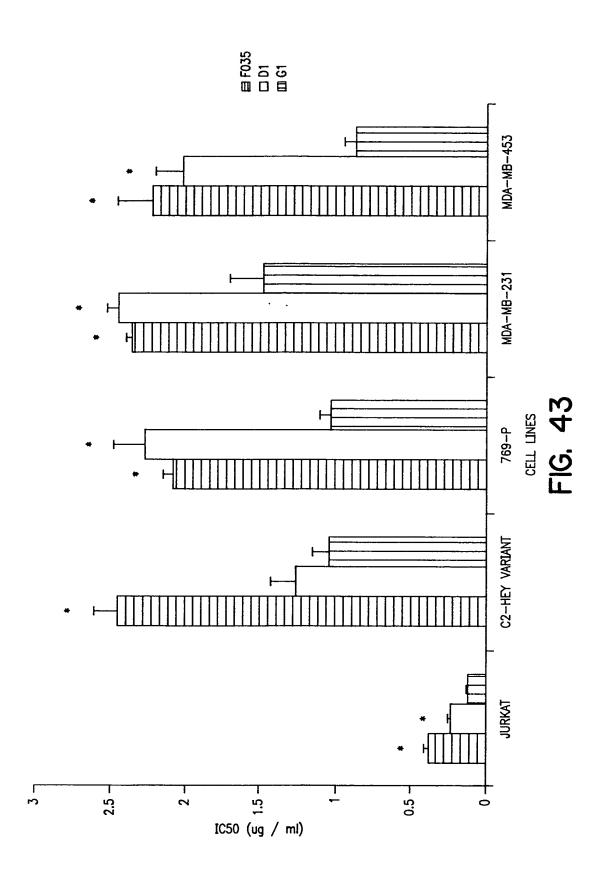
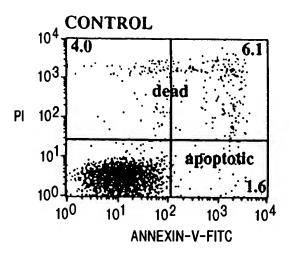


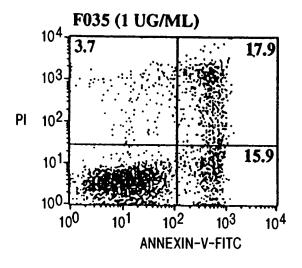
FIG. 42

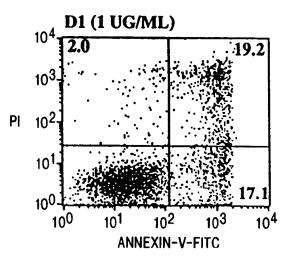


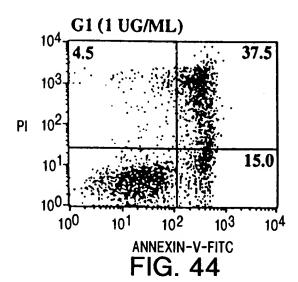


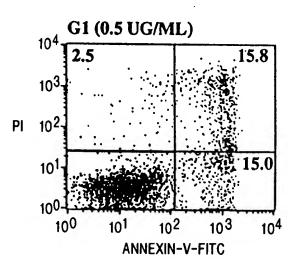














Fraction 35 treatment

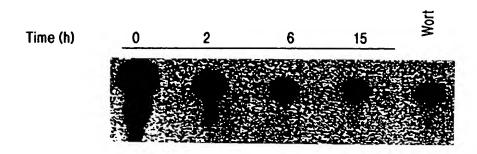


FIG. 45A

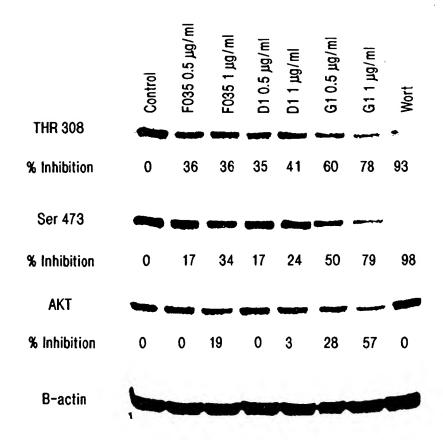


FIG. 45B



FIG. 46A

F035 (µg/ml) 0 0 1 1 2 2 4 4 0 0
TNF (100pM) - + - + - + - + - +
Wortmannin (1µM) - - - - - - - + +

Fold Activation 1.0 3.9 1.1 1.7 1.2 1.5 0.9 1.1 1.0 1.1

FIG. 46B

TNF
TNF
F035 + TNF
F035 + TNF
D1
D1
TNF
G1 + TNF
G1 + TNF

Fold Activation 1.0 3.4 0.9 1.2 0.9 0.9 0.9 1.2

NF-KB-

LPS 46C F035 + LPS Wortmannin + LPS

FIG. 46D

PHA + PMA
F035 + PHA + PMA
F035 + PHA + PMA



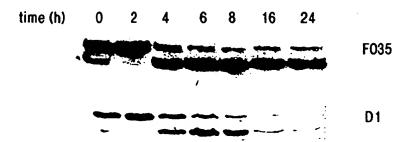


FIG. 47

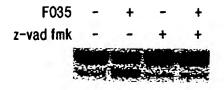
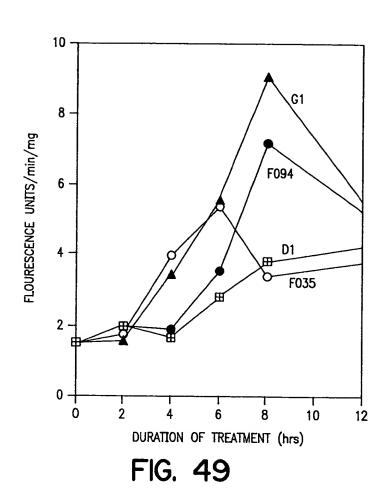


FIG. 48







time (h) 0 4 6 — Cytochrome c

FIG. 50



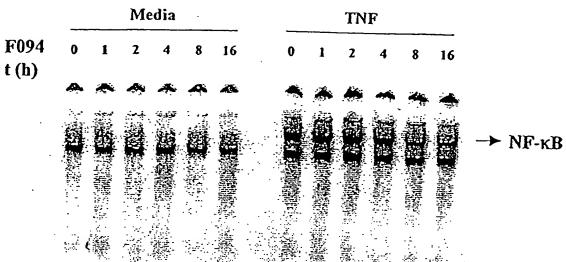


FIG. 51A

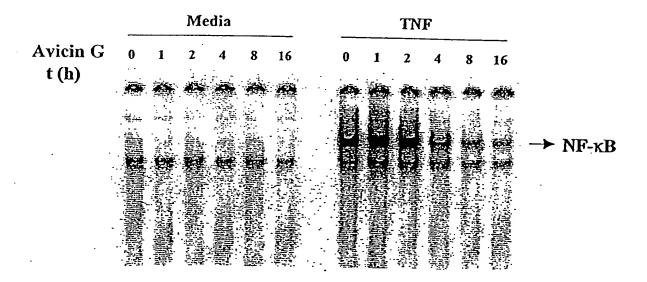


FIG. 51B



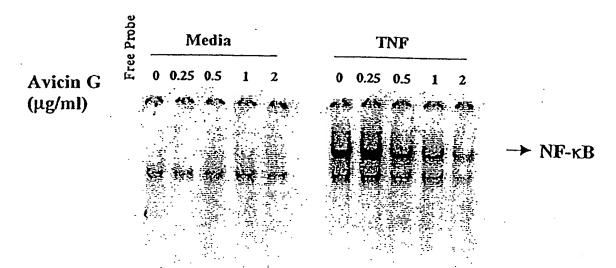


FIG. 52A

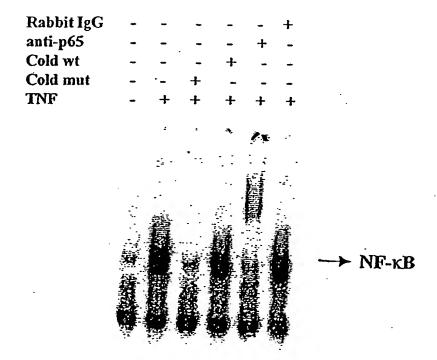


FIG. 52B



ΙκΒ

Time (min) 0 2 5 10 15 30

Untreated

Avicin G

FIG. 53A

p65

Time (min) 0 2 5 10 15 30

Untreated

Avicin G

FIG. 53B



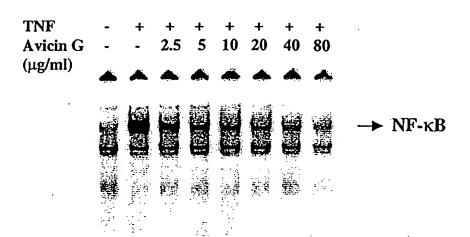


FIG. 54A

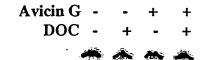




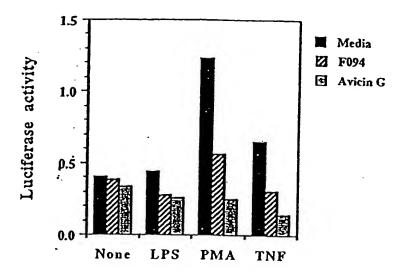
FIG. 54B

TNF - + - + + + + Avicin G - - + + - + DTT - - - + +



FIG. 54C





Inducers of NF-kB FIG. 55A

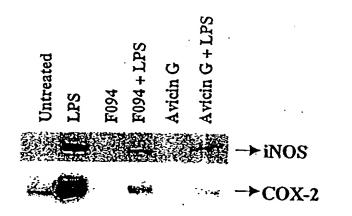
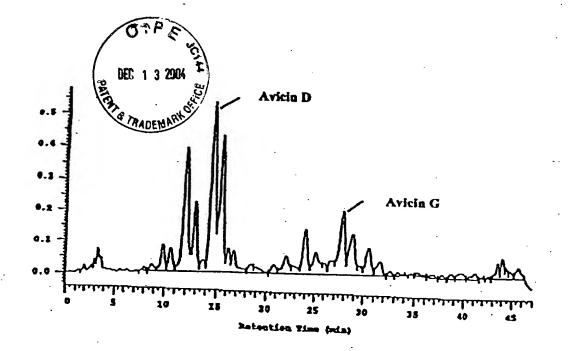


FIG. 55B



IC50 Values

HPLC Separation of the Avicins in F094

0.331-0.407 µg/ml

Avicin D

0.320-0.326 µg/ml

0.160-0.181 µg/ml

Avicin G



Annexin-V positive cells (%)

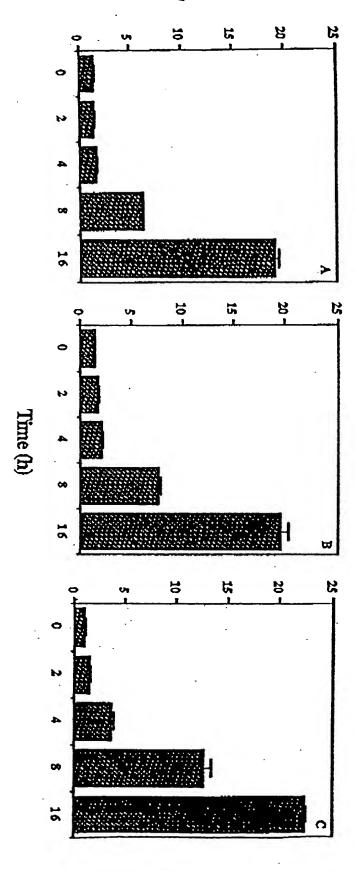


FIG. 57



Time (h) 0 0.5 1 2 4

Mixture

Fold increase 1.0 0.8 0.8 1.3 1.5

Avicin D

Fold increase 1.0 1.5 1.7 1.9 3.5

Avicin G



FIG. 59A Time (min) 0 1 2 5 10 20

∻ The Cyt-c

DEVD

2VAD-fmk

Avicia G

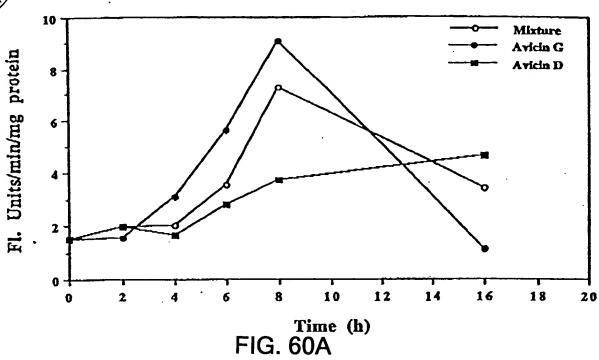
DEVD+Avicia G

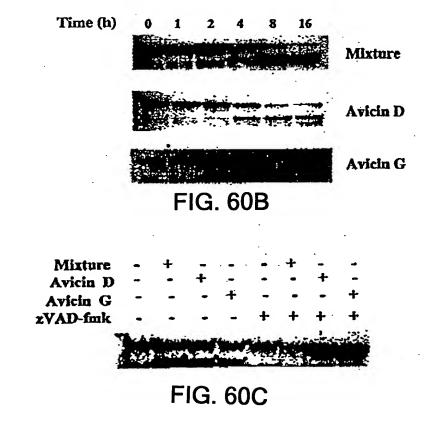
DEVD+Avicia G

DEVD-fmk+Avicia G

FIG. 59C









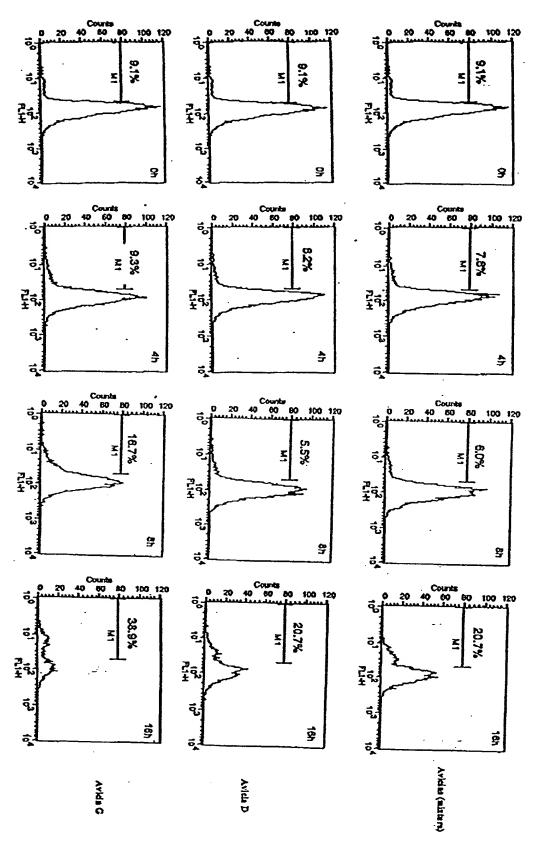


FIG. 61



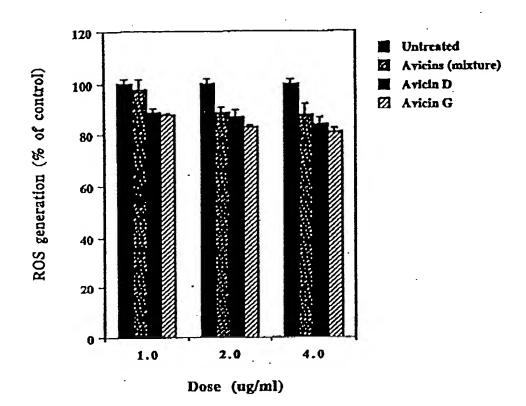


FIG. 62

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